# Plague Control in Northern Arizona

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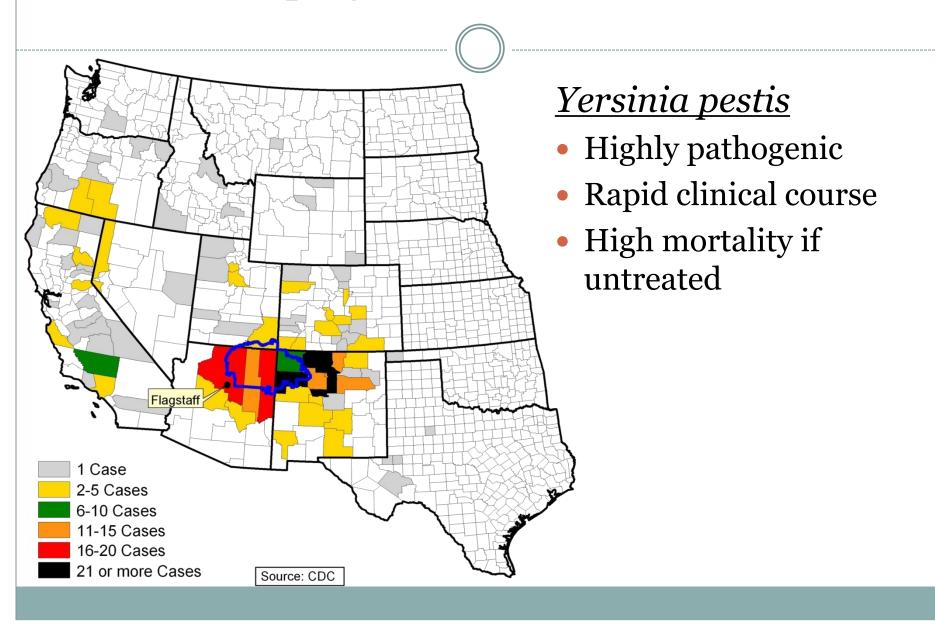


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NAU-MGGen (Center for Microbial Genetics and Genomics)
- BSL<sub>3</sub> Facility for bacterial pathogens

### Human plague in the US: 1970-2000



# Three historical plague pandemics

- 1st Pandemic: "Justinian plague"
  - o 6<sup>th</sup>-8<sup>th</sup> centuries AD
  - Affected North Africa, Europe, parts of Asia
  - Population losses estimated at 50-60%

#### • 2<sup>nd</sup> Pandemic:

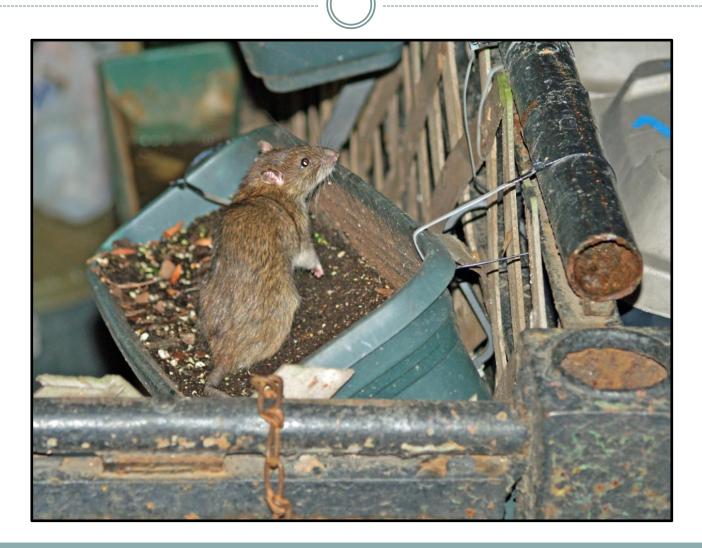
- o 14<sup>th</sup>-17<sup>th</sup> centuries AD
- Affected North Africa, Europe, parts of Asia
- Black Death epidemic: 1347-1351
  - x Est. 17-28 million killed, or ~30-40% of the European population

#### • 3<sup>rd</sup> Pandemic:

- o 19<sup>th</sup>-20<sup>th</sup> centuries AD
- o Global spread out of China: Africa, Americas, Australia, Europe
- Millions of deaths worldwide
- Control: improved hygiene (rat control) and antibiotics

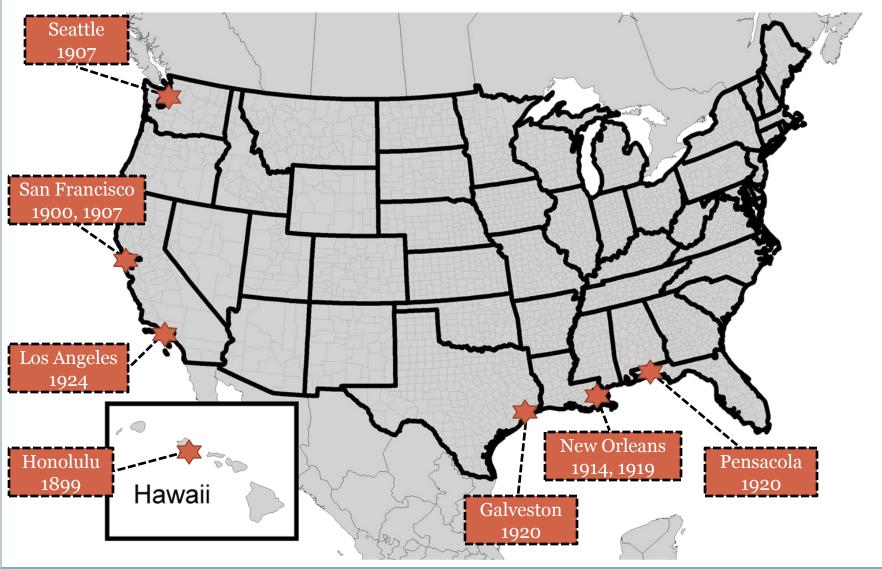
Perry and Fetherston. 1997. Clinical Microbiology Reviews 10:35-66.

Rats (*Rattus* spp.) and their fleas were important hosts during global spread and establishment of plague foci in cities



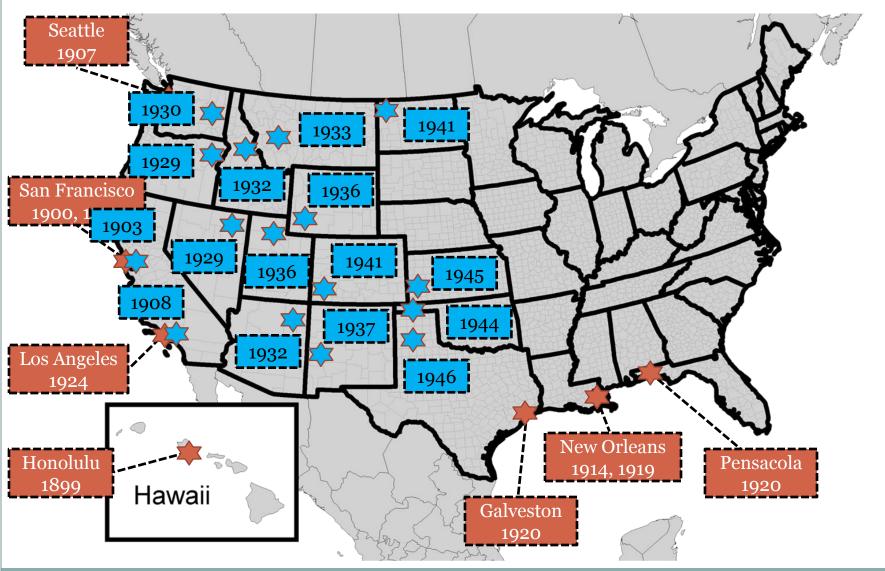
Rat in NYC, photo by David Shankbone

### Rat-borne Plague in Multiple Port Cities



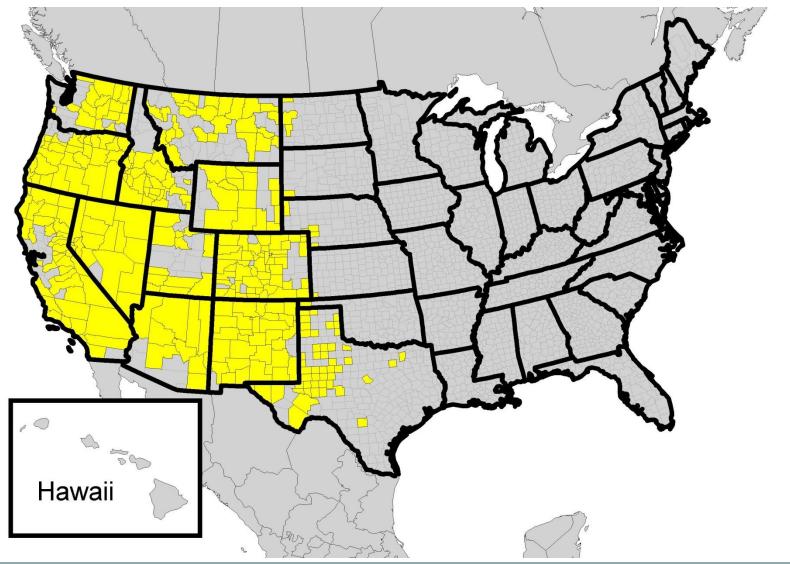
Eskey and Haas, 1940; Pollitzer 1951; Link 1955

#### Plague can establish in native rodents (esp. ground squirrels)



Eskey and Haas, 1940; Pollitzer 1951; Link 1955

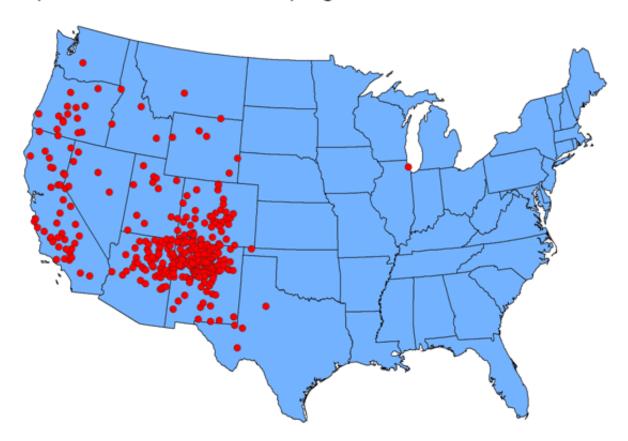
# Plague is Now Endemic in the Western US



CDC occurrence data for 1970-2000 as compiled by Cully and Williams 2001

# Recent Human Plague in the US

Reported cases of human plague--United States, 1970-2012



1 dot placed in county of exposure for each plague case

Source: www.cdc.gov/plague/maps/index.html

### Plague in humans

 Rare in US and difficult to diagnose without knowledge of risk factors

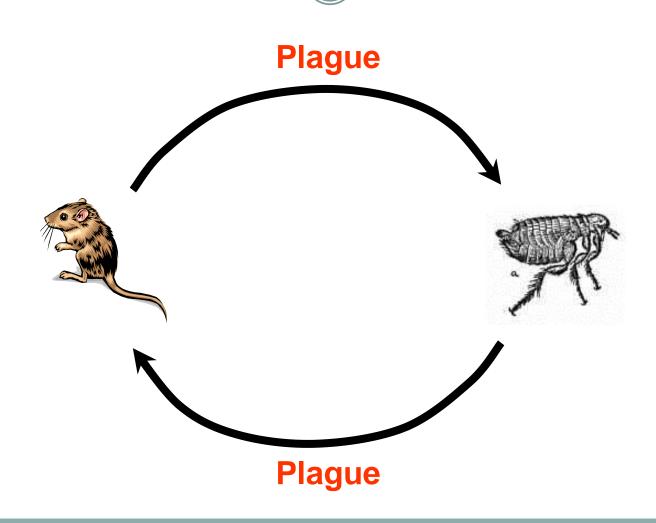
- Case example CO2012 (Fox news)
  - While camping in SW Colorado, girl handled a dead squirrel. Also had multiple insect bites.
  - o 107°F fever, "doctors were baffled by the cause"
  - Patient survived with antibiotics



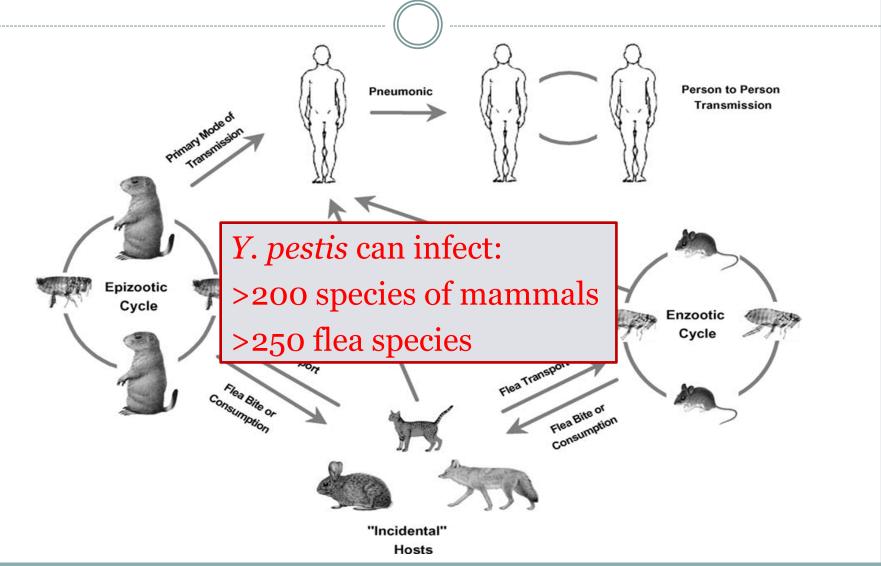
www.foxnews.com/health/2012/09/05/colorado-girl-recovering-from-bubonic-plague

- Antibiotic resistance is a concern
  - Y. pestis strain with resistance to multiple antibiotics found in Madagascar (Galimand et al. 2006; Welch et al. 2007)

### Basic Plague Ecology: a disease of rodents and fleas

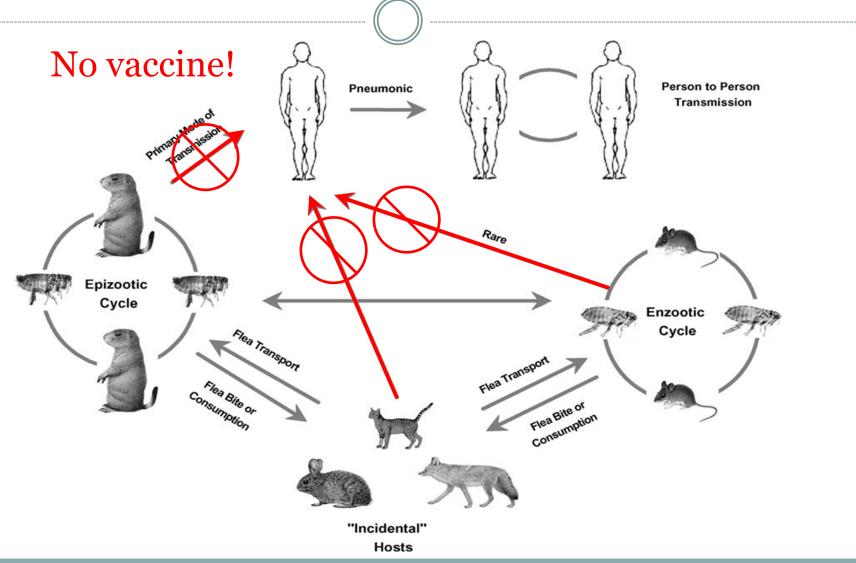


# More Complex Plague Ecology

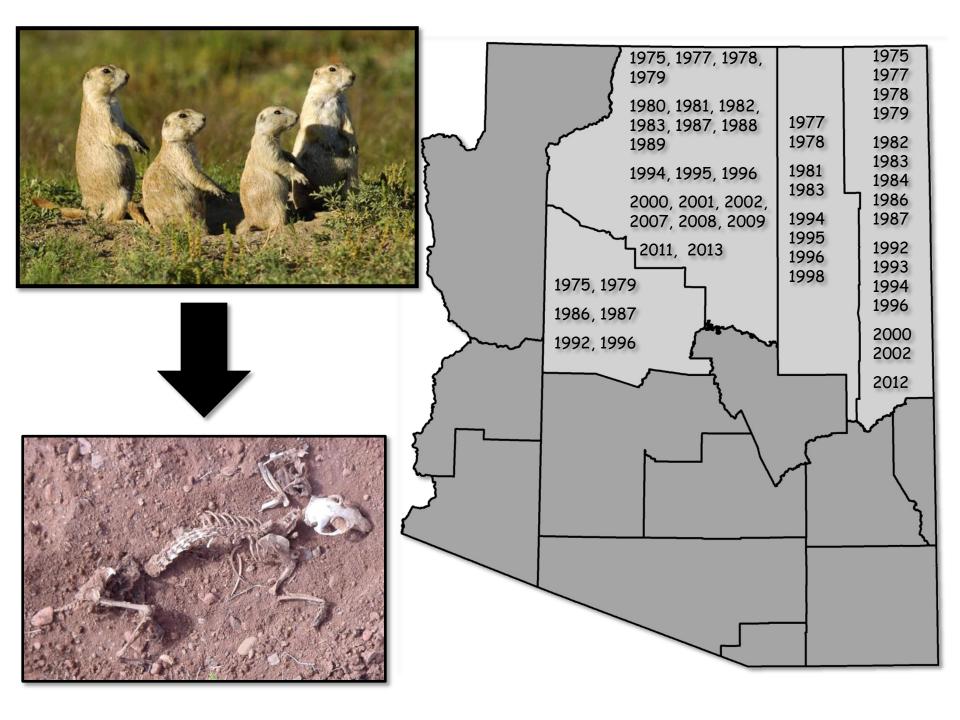


Gage and Kosoy. 2005. Annual Review of Entomology 50:505-528.

# More Complex Plague Ecology



Gage and Kosoy. 2005. Annual Review of Entomology 50:505-528.





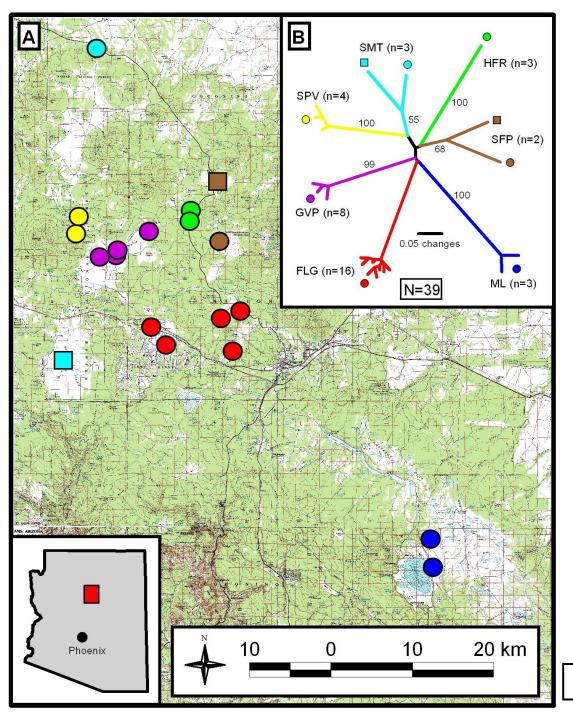
Sentinel species
- Monitored by CCHD





Photos: Katy Parise, Dave Wagner

See Girard and Wagner *et al.* 2004. PNAS 101:8408-8413.



#### Insights from genetics

- 2001 epizootic in Flagstaff area – sampled fleas from prairie dog burrows
- Plague is widespread in the environment
- Multiple introductions from plague reservoir

Fig. 2. Girard and Wagner et al. 2004

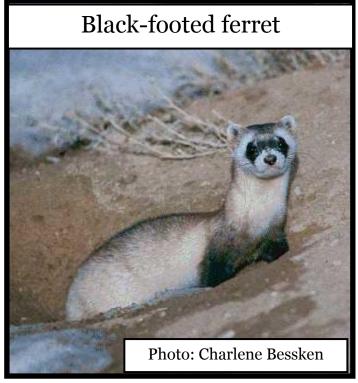
### Flea monitoring in northern AZ

- MGGen works with state and county health departments to screen fleas for *Y. pestis*.
  - DNA assay using PCR (polymerase chain reaction)
- Long-time collaboration with Coconino Co Health Dept.
  - Marlene Gaither, Hugh Murray, and team Thank You!
- Field sampling in short window before application of insecticide to burrows
  - NAU samples as many fleas as possible without interfering with fleat control

#### Flea control with Delta dust (deltamethrin)

- Dusting prairie dog burrows near human communities (Flagstaff)
- Flea control increases survival of 3 prairie dog species in plague-endemic areas (Biggins et al. 2010 Vector-Borne and Zoonotic Diseases)

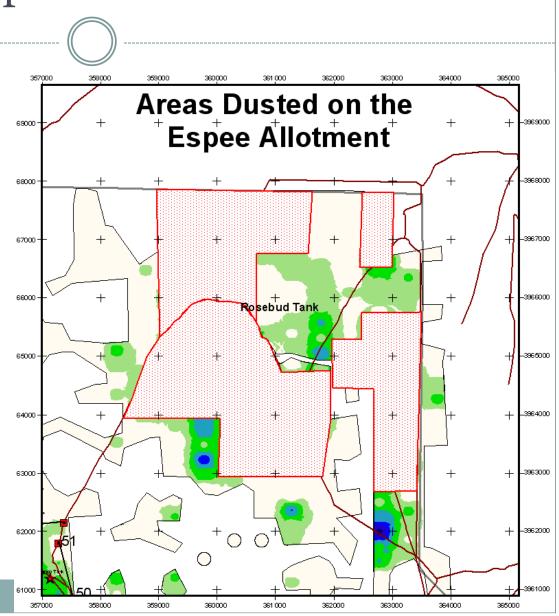




# Flea dusting at Espee Black-footed ferret site



- Successful insecticidal treatment in May-June 2009 (protected prairie dogs inside red areas)
- Non-treated areas (blue/green) inactive by August 2009
- We cultured Y. pestis from fleas sampled outside of protected areas



#### Alternative control method: Oral vaccines

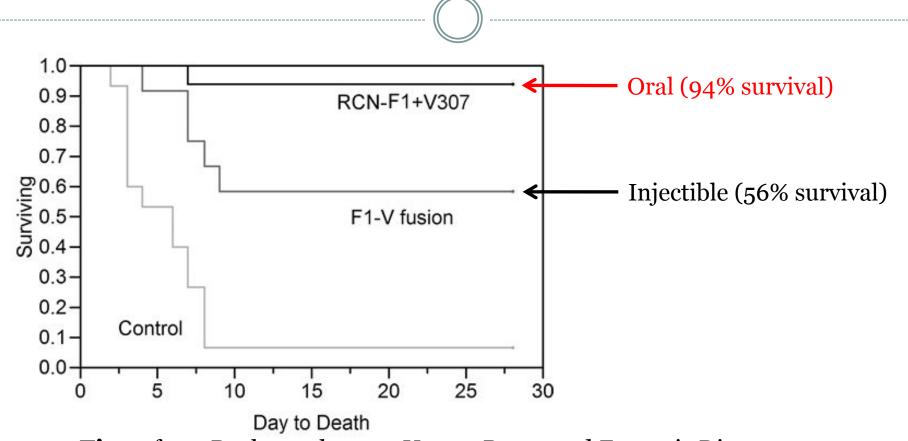


Fig. 2 from Rocke et al. 2010. Vector-Borne and Zoonotic Diseases.

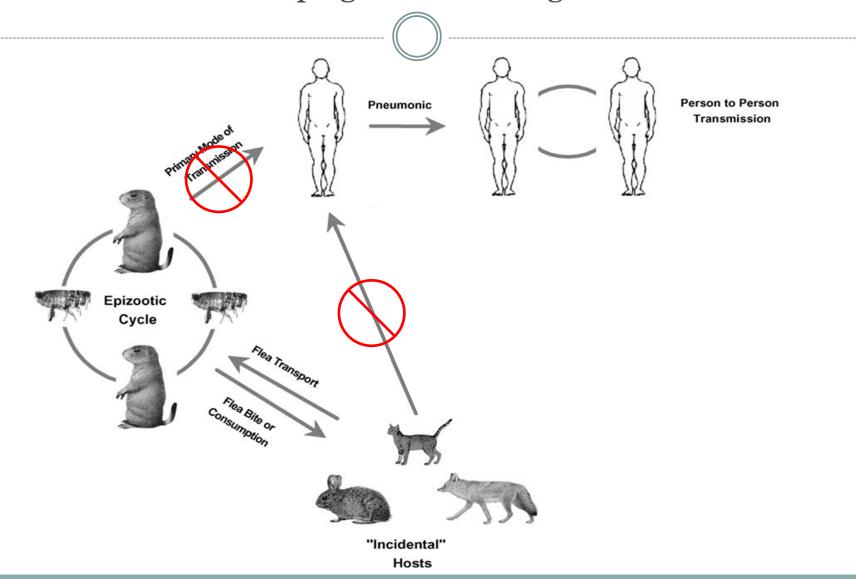
- Edible vaccine for *Y. pestis* now in field trials for prairie dogs (Tripp et al. 2014. J Wild Dis)

Map by Habitat Harmony, Inc. of Flagstaff

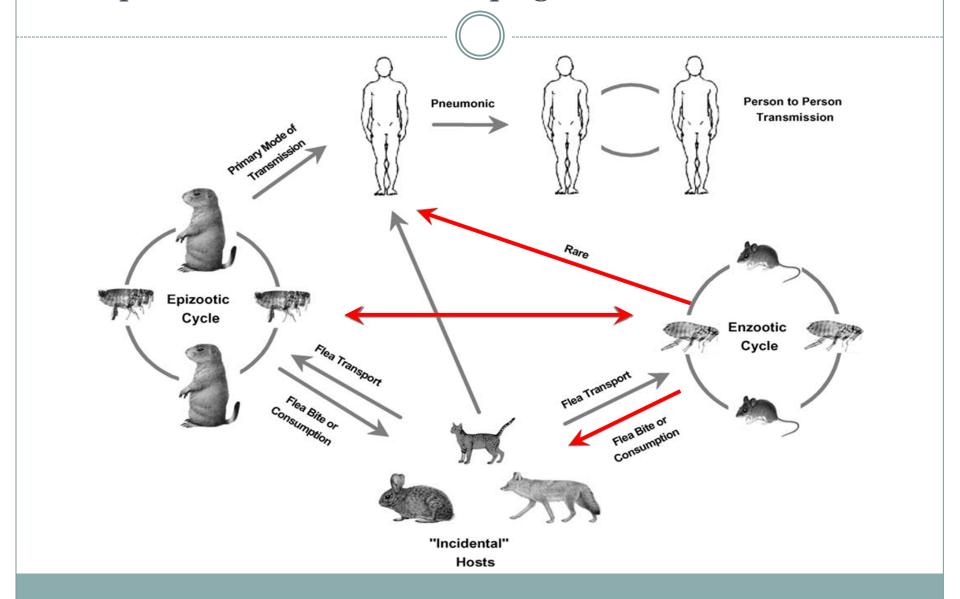
### Preventing transmission of *Y. pestis* to humans

- Use of sentinel species (prairie dogs) and fleas to detect plague
- Vector control (dusting for fleas)
- Education in rural and urban communities
  - Arizona Dept. of Health Services
     (http://www.azdhs.gov/phs/oids/vector/plague/index.htm)
  - Treat pets for fleas; understand the risk of pets acquiring plague or fleas from wild animals
  - Risk associated with human activity near rodent burrows & nests
  - Risk associated with handling/hunting wild animals (prairie dogs, coyotes, rabbits)

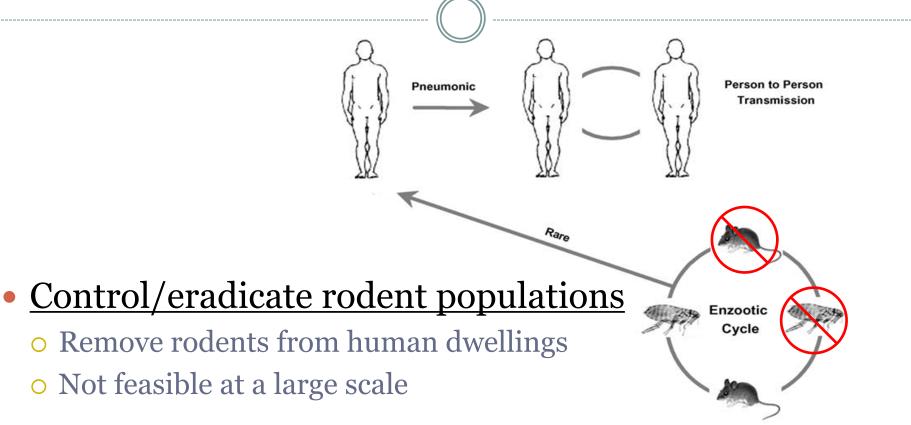
#### Current methods of plague control target incidental hosts



Is it possible to intervene with plague at its ultimate source?

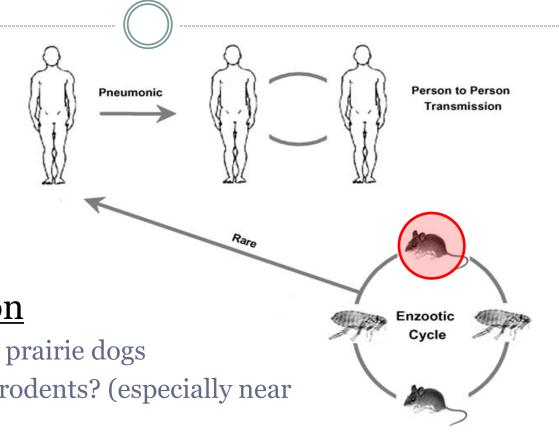


#### Is it possible to intervene with plague at its ultimate source?



- Control flea populations?
  - Not feasible for most rodent species
  - Frequent use could lead to resistant flea populations

#### Is it possible to intervene with plague at its ultimate source?



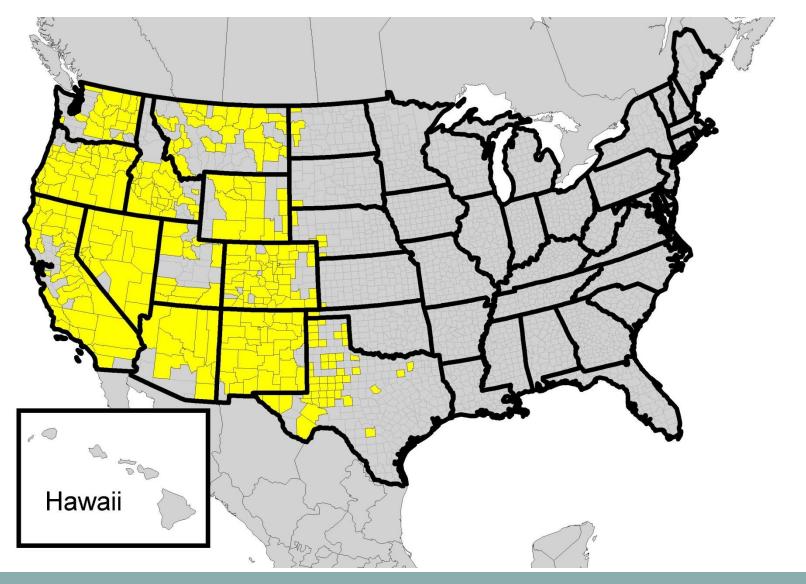
#### Rodent vaccination

- Oral vaccine tested in prairie dogs
- Possible to treat wild rodents? (especially near human dwellings)
- Our collaborators at ASU (Dr. Roy Curtiss and Dr. Wei Sun) are developing an oral plague vaccine for laboratory mice

# Vaccination would be targeted at a local scale (not regional) - Short timeframes



#### Plague likely to remain endemic, but it is possible to reduce the threat of transmission to humans



### Conclusions and Take Home

- Humans have dispersed *Y. pestis* globally, and it is now endemic in the western U.S.
- Y. pestis cycles in native flea and rodent reservoirs
- Currently, plague control targets incidental hosts
  - Near human dwellings
  - Black-footed ferret sites
- Flea treatment is a widely used control method
- Intervention of plague at its ultimate source would require an alternative strategy:
  - o Vaccinating small rodents (possible?)

### Acknowledgements



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